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                 FSTA enhanced with new thesaurus edition
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      3
         AUG 06
NEWS
         AUG 13
                 CA/CAplus enhanced with additional kind codes for granted
         AUG 20
                 CA/CAplus enhanced with CAS indexing in pre-1907 records
NEWS
      5
                 Full-text patent databases enhanced with predefined
NEWS
      6
         AUG 27
                 patent family display formats from INPADOCDB
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      7
         AUG 27
                 USPATOLD now available on STN
                 CAS REGISTRY enhanced with additional experimental
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      8
         AUG 28
                 spectral property data
                 STN AnaVist, Version 2.0, now available with Derwent
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         SEP 07
                 World Patents Index
                 FORIS renamed to SOFIS
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         SEP 13
                 INPADOCDB enhanced with monthly SDI frequency
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         SEP 13
                 CA/CAplus enhanced with printed CA page images from
NEWS 12
         SEP 17
                 1967-1998
                 CAplus coverage extended to include traditional medicine
NEWS 13
         SEP 17
                 patents
                 EMBASE, EMBAL, and LEMBASE reloaded with enhancements
NEWS 14 SEP 24
        OCT 02
                 CA/CAplus enhanced with pre-1907 records from Chemisches
NEWS 15
                 Zentralblatt
NEWS 16 OCT 19
                 BEILSTEIN updated with new compounds
                 Derwent Indian patent publication number format enhanced
        NOV 15
NEWS 17
NEWS 18 NOV 19
                 WPIX enhanced with XML display format
NEWS 19 NOV 30
                 ICSD reloaded with enhancements
                 LINPADOCDB now available on STN
NEWS 20 DEC 04
NEWS 21 DEC 14
                 BEILSTEIN pricing structure to change
                 USPATOLD added to additional database clusters
NEWS 22 DEC 17
                 IMSDRUGCONF removed from database clusters and STN
NEWS 23 DEC 17
                 DGENE now includes more than 10 million sequences
NEWS 24
        DEC 17
                 TOXCENTER enhanced with 2008 MeSH vocabulary in
NEWS 25
        DEC 17
                 MEDLINE segment
                 MEDLINE and LMEDLINE updated with 2008 MeSH vocabulary
NEWS 26
        DEC 17
                 CA/CAplus enhanced with new custom IPC display formats
NEWS 27
        DEC 17
NEWS 28
        DEC 17
                 STN Viewer enhanced with full-text patent content
                 from USPATOLD
                 STN pricing information for 2008 now available
NEWS 29
         JAN 02
                 CAS patent coverage enhanced to include exemplified
         JAN 16
NEWS 30
                 prophetic substances
NEWS EXPRESS 19 SEPTEMBER 2007: CURRENT WINDOWS VERSION IS V8.2,
              CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
              AND CURRENT DISCOVER FILE IS DATED 19 SEPTEMBER 2007.
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              Welcome Banner and News Items
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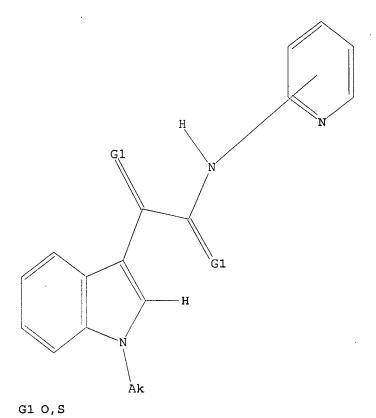
=>
Uploading C:\Program Files\Stnexp\Queries\10686809 NEW_1242008.str

L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> s 11 full

FULL SEARCH INITIATED 15:23:48 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 2261 TO ITERATE

100.0% PROCESSED 2261 ITERATIONS

143 ANSWERS

TOTAL

SINCE FILE

SEARCH TIME: 00.00.01

L2 143 SEA SSS FUL L1

=> file medline caplus wpids uspatfull

COST IN U.S. DOLLARS

FULL ESTIMATED COST ENTRY SESSION 178.36 178.57

FILE 'MEDLINE' ENTERED AT 15:23:58 ON 24 JAN 2008

FILE 'CAPLUS' ENTERED AT 15:23:58 ON 24 JAN 2008
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=> s 12

SAMPLE SEARCH INITIATED 15:24:02 FILE 'WPIDS'
SAMPLE SCREEN SEARCH COMPLETED - 43 TO ITERATE

100.0% PROCESSED 43 ITERATIONS

SEARCH TIME: 00.00.02

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 234 TO 626 PROJECTED ANSWERS: 9 TO 179

L3 185 L2

=> s 13 and (cancer? or tumor?)

L4 58 L3 AND (CANCER? OR TUMOR?)

=> s 14 and angiogenesis

L5 13 L4 AND ANGIOGENESIS

=> s 14 and (multidrug or multi-drug)

L6 15 L4 AND (MULTIDRUG OR MULTI-DRUG)

=> d 16 1-15 ibib, abs, hitstr

L6 ANSWER 1 OF 15 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2003:757332 CAPLUS

DOCUMENT NUMBER: 139:276902

TITLE: Preparation of 2-(3-indoly1)-2-oxoacetamide

derivatives as angiogenesis inhibitors and anticancer

agents

INVENTOR(S): Chen, Chiung-tong; Chen, Shu-jen; Hsu, Ming-chu;

Hwang, Der-ren; Li, Wen-tai; Lin, Chu-chung

PATENT ASSIGNEE(S): National Health Research Institutes, Taiwan

SOURCE: U.S. Pat. Appl. Publ., 26 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| | PATENT NO. | KIND | DATE | APPLICATION NO. DATE | |
|---|------------------------|--------|------------|--------------------------|-----|
| | | | | | |
| | US 2003181482 | A1 | 20030925 | US 2002-310711 200212 | 205 |
| | US 6903104 | B2 | 20050607 | | |
| | US 2005234098 | A1 | 20051020 | US 2005-145628 200506 | 506 |
| F | PRIORITY APPLN. INFO.: | | | US 2001-337962P P 200112 | 206 |
| | | | | US 2002-310711 A1 200212 | 205 |
| | THER SOURCE(S): | MARPAT | 139:276902 | | |

This invention relates to novel heteroatom containing compds. [R1 = AB independently each (un) substituted isoxazolyl, thiazolyl, isothiazolyl, 1,3,4-thiadiazolyl, 1,3-benzothiazolyl, quinolyl, isoquinolyl, thionaphthenyl, or benzofuranyl; R2 = independently H, each (un) substituted C1-10 alkyl or aryl; or R1 and R2 are taken together with the nitrogen atom to which they are attached to form an (un) substituted 5-8 membered ring comprising C, N, S, or O atoms but not to form 4-phenylpiperazin-1-yl, 4-(pyridin-4-yl)piperazin-1-yl, 4-(pyridin-2-yl)piperazin-1-yl, 4-(2-nitrophenyl)piperazin-1-yl, 4-(3,5-dimethoxyphenyl)piperazin-1-yl, or 4-[bis(4fluorophenyl)methyl]piperazin-1-yl; R3 = independently each (un) substituted C1-10 alkyl, C2-10 alkenyl, C2-10 alkynyl, C3-10 cycloalkyl, C4-10 cycloalkenyl, aryl, heteroaryl, or heterocyclyl; R4 = each independently H, NO2, halo, cyano, R7, OR7, CO2R7, SR7, NR7R7, C(O)R7, C(O)R7, OC(O)R7, S(O)2R7, S(O)2R7, NR7C(O)R7, NR7C(O)R7, NR7(CO2R7), NR7S(O)2NR7R7, or NR7S(O)2R7, S(O)2OR7; n = 0, 1, 2, 3, or 4; R7 = independently H, each (un) substituted C1-10 alkyl, C2-10 alkenyl C2-10 alkynyl, C3-10 cycloalkyl, aryl, heteroaryl, or heterocyclyl]. These compds. have potent anticancer, cytotoxic, and anti-angiogenic activity and are useful for the prevention and treatment of diseases, in particular a cancer including a human leukemia, sarcoma, osteosarcoma, lymphoma, melanoma, ovarian, skin, testicular, gastric, pancreatic, renal, breast, prostate colorectal, head and neck, brain, esophageal, bladder, adrenal cortical, lung, bronchus, endometrial, cervical or hepatic cancer, or cancer of unknown primary site. Moreover the cancer is a drug resistance phenotype of which the cancer cells express P-glycoprotein (MDR), multidrug resistance-associated proteins (MRP), lung cancer resistance- associated proteins (LRP), breast cancer resistance proteins (BCRP) or other proteins associated with resistance to anticancer drugs. Thus, a solution of 1.17 g indole 10 mL THF was added dropwise to a suspension of 1.34 g potassium tert-butoxide in 10 mL THF, stirred at room temperature for 2 h, then treated dropwise with a solution of 1.32 g 5-(chloromethyl)-3-methylisoxazole in 5 mL THF, and allowed stand for 4 h, and quenched by adding 10 mL saturated ammonium chloride to give, after workup and silica gel chromatog., 1.61 g 5-(1H-1-indolylmethyl)-3methylisoxazole (II) (76%). A solution of 212 mg II in 10 mL di-Et ether was added to 254 mg oxalyl chloride dropwise at 0°, stirred at 0° for 3 h, evaporated to remove the solvent, dissolved in 5 mL THF, treated with a solution of 114 mg 3-methyl-5-isothiazolamine and 1 mL Et3N in 10 mL THF dropwise, stirred for 10 h, and then treated with 1 N NaOH (4 mL) to give, after workup and crystallization, 0.27 g I (R1 = 3-methyl-5isothiazolyl, R2 = R4 = H, R3 = 3-methyl-5-isoxazolyl) (III) (71%). III in vitro inhibited the growth of human cancer cell lines DLD1, HA-22T, HEP G2, HONE1, HR, and NUGC3 with IC50 of 41, 123, 93, 4, 8, and 12 nM, resp. 501921-65-9P, N-(4-Pyridyl)-2-[1-(4-cyanobenzyl)-1H-indol-3-yl]-2-IT oxoacetamide RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (preparation of (3-indoly1)oxoacetamide derivs. as angiogenesis inhibitors

1H-Indole-3-acetamide, 1-[(4-cyanophenyl)methyl]- α -oxo-N-4-pyridinyl-

and anticancer agents)

501921-65-9 CAPLUS

(CA INDEX NAME)

RN

CN

REFERENCE COUNT:

THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS 21 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 2 OF 15 CAPLUS COPYRIGHT 2008 ACS on STN L6

ACCESSION NUMBER:

2003:235682 CAPLUS

DOCUMENT NUMBER:

138:378576

TITLE:

Synthesis and Biological Evaluation of N-Heterocyclic Indolyl Glyoxylamides as Orally Active Anticancer

Agents

AUTHOR(S):

Li, Wen-Tai; Hwang, Der-Ren; Chen, Ching-Ping; Shen, Chien-Wei; Huang, Chen-Long; Chen, Tung-Wei; Lin, Chi-Hung; Chang, Yee-Ling; Chang, Ying-Ying; Lo, Yue-Kan; Tseng, Huan-Yi; Lin, Chu-Chung; Song,

Jeng-Shin; Chen, Hua-Chien; Chen, Shu-Jen; Wu, Se-Hui;

Chen, Chiung-Tong

CORPORATE SOURCE:

Division of Biotechnology and Pharmaceutical Research,

National Health Research Institutes, Taipei, 114,

Taiwan

SOURCE:

Journal of Medicinal Chemistry (2003), 46(9),

1706-1715

CODEN: JMCMAR; ISSN: 0022-2623

PUBLISHER:

American Chemical Society

DOCUMENT TYPE:

Journal

LANGUAGE:

English

OTHER SOURCE(S):

CASREACT 138:378576

A series of N-heterocyclic indolyl glyoxylamides were synthesized and evaluated for in vitro and in vivo anticancer activities. They exhibited a broad spectrum of anticancer activity not only in murine leukemic cancer cells but also in human gastric, breast, and uterus cancer cells as well as their multidrug resistant sublines with a wide range of IC50 values. They also induced apoptosis and caused DNA fragmentation in human gastric cancer cells. Among the compds. studied, N1-(3-Methyl-5-isothiazolyl)-2-1-[(3-methyl-5isoxazolyl)methyl]-1H-3-indolyl-2-oxoacetamide (I) showed the most potent activity of growth inhibition (IC50 = 17-1711 nM) in several human cancer cells. Given orally, compds. I and N1-(3-Methyl-5isothiazoly1) -2-[1-(4-cyanobenzy1)-1H-3-indoly1]-2-oxoacetamide dose-dependently prolonged the survival of animals inoculated with P388 leukemic cancer cells. N-Heterocyclic indolyl glyoxylamides may be useful as orally active chemotherapeutic agents against cancer and refractory cancerous diseases of multidrug resistance phenotype.

528593-64-8P TT

RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(synthesis and biol. evaluation of N-Heterocyclic indolyl glyoxylamides as orally active anticancer agents in relation to apoptosis induction and partition coefficient)

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 3 OF 15 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER:

2003:221515 CAPLUS

DOCUMENT NUMBER:

138:238008

TITLE:

Preparation of 3-glyoxlylamide indoles as anticancer

agents useful against multidrug-resistant

cancer cells

INVENTOR(S):

Koya, Keizo; Sun, Lijun; Ono, Mitsunori; Liang, Guiqing; James, David; Li, Hao; Xia, Zhi-Qiang

PATENT ASSIGNEE(S):

SBR Pharmaceuticals Corp., USA

SOURCE:

PCT Int. Appl., 55 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | | | | | DATE | | APPLICATION NO. | | | | | | | | | | |
|------------|------------------------|-----|-------------------------|----------------------|-------------------------|-----|-----------------|----------|----------|----------------|-------|-----|-----|------|-----|------|-----|
| | | | | | | | | | | | | | | | | | |
| | | | | | | | WO 2002-US27513 | | | | | | 21 | 1020 | 828 | | |
| WO | 2003022280 A3 20030522 | | | | | | | | | | | | | | | | |
| | W: | ΑE, | AG, | AL, | AM, | AT, | AU, | ΑZ, | ВA, | BB, | BG, | BR, | BY, | ΒŹ, | CA, | CH, | CN, |
| | | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EC, | EE, | ES, | FI, | GB, | GD, | GE, | GH, |
| | | GM, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | KΕ, | KG, | ΚP, | KR, | ΚZ, | LC, | LK, | LR, |
| | | LS, | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | ΜZ, | NO, | ΝZ, | OM, | PH, |
| | | PL, | PT, | RO, | RU, | SD, | SE, | SG, | SI; | SK, | SL, | ΤĴ, | TM, | TN, | TR, | TT, | TZ, |
| | | | | | | | VN, | | | | | | | | * | | |
| | RW: | GH, | GM, | KE, | LS, | MW | MZ, | SD, | SL, | SZ, | TZ, | UG, | ZM, | ZW, | AM, | AZ, | BY, |
| | | KG, | KZ, | MD, | RU, | TJ | TM, | AT, | BE, | BG, | CH, | CY, | CZ, | DE, | DK, | EE, | ES, |
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| CA | 2460 | | • | • | • | | | | • | | • | | | | 2 | 0020 | 328 |
| | | | | | | | | | | AU 2002-323474 | | | | | | | |
| | | | 20040616 EP 2002-757457 | | | | 00208 | | | | | | | | | | |
| | | | | | | | ES, | | | | | | | | | | |
| | | | | | | | RO, | | | | | | | | | , | , |
| .тъ | 2005 | | | | | | | | | | | | | | | 0020 | 328 |
| | 2005504790 T | | | 30515 US 2002-232394 | | | | | | | | | | | | | |
| | | | | | | | | | 20020025 | | , 2 , | | | | | | |
| | | | | | | | | | | 20050524 | | | | | | | |
| US | US 2006004044 | | AΙ | | 20060105 US 2005-136074 | | | 20050524 | | | | | | | | | |

PRIORITY APPLN. INFO.:

US 2001-322022P WO 2002-US27513 P 20010913

US 2002-232394

W 20020828 A1 20020829

OTHER SOURCE(S):

MARPAT 138:238008

GI

The anti-cancer compound has a structural formula I wherein Z1 and Z2 are independently O, S, NOR5 or NR5, and R1-R5 are H, aliphatic group, aryl group or other specifically defined groups. Thus, 2 - (1 - (4 - chloro-benzyl) - 1 - indo - 3 - yl) - N - (3 - methyl-isothiazol - 5 - yl) - 2 - oxo-acetamide was prepared from oxylyl chloride 5.1 mmol, $1 - (4 \cdot \text{chlorobenzyl}) - \text{indole}$ (4.14 mmol) and 5-amino-3-methylisothiazole (9.73 mmol), and demonstrated significantly high anti-cancer activity (IC50 0.0005 μ M) against five cancer lines with wide variety of multidrug-resistant cancer cell types (MDA 435, HL 60, DU 146, MES SA, and H2).

IT 501921-60-4P 501921-65-9P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of glyoxlylamide indoles as anticancer agents useful against multidrug-resistant cancer cells)

RN 501921-60-4 CAPLUS

CN 1H-Indole-3-acetamide, N-[5-(aminocarbonyl)-2-pyridinyl]-1-[(4-chlorophenyl)methyl]-α-oxo- (CA INDEX NAME)

RN 501921-65-9 CAPLUS

CN 1H-Indole-3-acetamide, 1-[(4-cyanophenyl)methyl]- α -oxo-N-4-pyridinyl-(CA INDEX NAME)

L6 ANSWER 4 OF 15 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2001:915607 CAPLUS

DOCUMENT NUMBER: 136:193482

TITLE: New small-molecule tubulin inhibitors

AUTHOR(S): Bacher, G.; Beckers, T.; Emig, P.; Klenner, T.;

Kutschert, B.; Nickel, B.

CORPORATE SOURCE: IUPAC Commission, Research & Development Oncology,

ASTA Medica AG, Frankfurt, 60314, Germany

SOURCE: Pure and Applied Chemistry (2001), 73(9), 1459-1464

CODEN: PACHAS; ISSN: 0033-4545

PUBLISHER: International Union of Pure and Applied Chemistry

DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

AB A review. The variety of biol. agents directed toward the tubulin system exceeds those acting on DNA, making it an important target for cancer chemotherapy. However, the complicated chemical structures and restricted access to the natural resources, in combination with the development of drug resistance, limit the first generation of natural products. Considerable efforts in the search and synthesis of new synthetic compds., such as small mol. tubulin inhibitors, gave access to novel potential/promising drugs. Among these substances, two series of novel, easily accessible indole classes were identified as tubulin-destabilizing agents. Owing to the synthetic nature, potent in vitro and in vivo antitumoral activity, and efficacy against multidrug-resistant (MDR) tumors, D-24851 and D-64131 have significant potential in cancer treatment.

IT 204205-90-3, D-24851

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (small-mol. tubulin inhibitors)

RN 204205-90-3 CAPLUS

CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 5 OF 15 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2001:58000 CAPLUS

DOCUMENT NUMBER: 134:290069

TITLE: D-24851, a novel synthetic microtubule inhibitor,

exerts curative antitumoral activity in vivo, shows

efficacy toward multidrug-resistant tumor cells, and lacks neurotoxicity

AUTHOR(S): Bacher, Gerald; Nickel, Bernd; Emig, Peter; Vanhoefer,

Udo; Seeber, Siegfried; Shandra, Alexei; Klenner,

Thomas; Beckers, Thomas

CORPORATE SOURCE: Department of Cancer Research, ASTA Medica AG,

Frankfurt am Main, 60314, Germany

SOURCE: Cancer Research (2001), 61(1), 392-399

CODEN: CNREA8; ISSN: 0008-5472

PUBLISHER: American Association for Cancer Research

DOCUMENT TYPE: Journal LANGUAGE: English

N-(pyridin-4-yl)-[1-(4-chlorbenzyl)indol-3-yl]glyoxylamide (D-24851) is a novel synthetic compound that was identified in a cell-based screening assay to discover cytotoxic drugs. D-24851 destabilizes microtubules and blocks cell cycle transition specifically at G2-M phase. The binding site of D-24851 does not overlap with the tubulin binding sites of known microtubule-destabilizing agents like vincristine or colchicine. vitro, D-24851 has potent cytotoxic activity toward a panel of established human tumor cell lines including SKOV3 ovarian cancer, U87 glioblastoma, and ASPC-1 pancreatic cancer cells. In vivo, oral D-24851 treatment induced complete tumor regressions (cures) in rats bearing Yoshida AH13 sarcomas. Of importance is that the administration of curative doses of D-24851 to the animals revealed no systemic toxicity in terms of body weight loss and neurotoxicity in contrast to the administration of paclitaxel or vincristine. Interestingly, multidrug-resistant cell lines generated by vincristine-driven selection or transfection with the Mr 170,000 P-glycoprotein encoding cDNA were rendered resistant toward paclitaxel, vincristine, or doxorubicin but not towards D-24851 when compared with the parental cells. Because of its synthetic nature, its oral applicability, its potent in vitro and in vivo antitumoral activity, its efficacy against multidrug-resistant tumors, and the lack of neurotoxicity, D-24851 may have significant potential for the treatment of various malignancies.

IT 204205-90-3, D 24851

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(D-24851, a novel synthetic microtubule inhibitor, exerts curative antitumoral activity in vivo, shows efficacy toward multidrug -resistant tumor cells, and lacks neurotoxicity)

RN 204205-90-3 CAPLUS

CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 6 OF 15 USPATFULL on STN

ACCESSION NUMBER:

2007:169455 USPATFULL

TITLE:

Combination bacteriolytic therapy for the treatment of

tumors

36

INVENTOR(S):

Dang, Long, Baltimore, MD, UNITED STATES Bettegowda, Chetan, Baltimore, MD, UNITED STATES Kenzler, Kenneth W., Bel Air, MD, UNITED STATES

Vogelstein, Bert, Baltimore, MD, UNITED STATES

PATENT ASSIGNEE(S):

The Johns Hopkins University, Baltimore, MD, UNITED

STATES, 21218 (U.S. corporation)

| | NUMBER | KIND | DATE | |
|---------------------|-----------------|------|----------|--------------|
| • | | | | |
| PATENT INFORMATION: | US 2007148135 | A1 | 20070628 | |
| APPLICATION INFO.: | US 2004-568765 | A1 | 20041021 | (10) |
| | WO 2004-US34625 | | 20041021 | |
| | | | 20070212 | PCT 371 date |

NUMBER DATE

PRIORITY INFORMATION:

US 2003-512923P 20031022 (60)

DOCUMENT TYPE:

Utility

24

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

BANNER & WITCOFF, LTD., 1100 13th STREET, N.W., SUITE

1200, WASHINGTON, DC, 20005-4051, US

NUMBER OF CLAIMS:

EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 16 Drawing Page(s)

LINE COUNT: 1016

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Current approaches for treating cancer are limited, in part, by the inability of drugs to affect the poorly vascularized regions of tumors. We have found that spores of anaerobic bacteria in combination with agents which interact with microtubules can cause the destruction of both the vascular and avascular compartments of tumors. Two classes of microtubule inhibitors were found to exert markedly different effects. Some agents that inhibited microtubule synthesis, such as vinorelbine, caused rapid, massive hemorrhagic necrosis when used in combination with spores. In contrast, agents that stabilized microtubules, such as the taxane docetaxel, resulted in slow tumor regressions that killed most neoplastic cells. Remaining cells in the poorly perfused regions of tumors could be eradicated by sponzlated bacteria. Mechanistic studies showed that the microtubule destabilizers, but not the microtubule stabilizers, radically reduced blood flow to tumors, thereby enlarging the hypoxic niche in which spores could germinate. A single intravenous

injection of spores plus selected microtubule-interacting agents was able to cause regressions of several tumors in the absence of excessive toxicity.

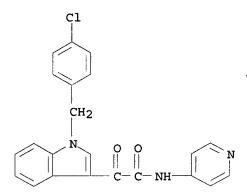
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

204205-90-3, D-24851

(combination bacteriolytic therapy for the treatment of tumors using spores of anaerobic bacteria and microtubule agents)

RN 204205-90-3 USPATFULL

1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-CN pyridinyl- (CA INDEX NAME) .



ANSWER 7 OF 15 USPATFULL on STN L6

ACCESSION NUMBER:

2006:167882 USPATFULL

TITLE:

Bis(thio-hydrazide amides) for treatment of hyperplasia

INVENTOR(S):

Sherman, Matthew L., Newton, MA, UNITED STATES Vaghefi, Farid, Burlington, MA, UNITED STATES Chen, Lan Bo, Lexington, MA, UNITED STATES

| | NUMBER | KIND | DATE | |
|--------------|----------------|------|----------|------|
| | | | | |
| INFORMATION: | US 2006142393 | A1 | 20060629 | |
| ATION INFO.: | US 2005-226929 | A1 | 20050914 | (11) |

PATENT APPLICA

> DATE NUMBER

PRIORITY INFORMATION:

US 2004-610270P 20040916 (60)

DOCUMENT TYPE: Utility

FILE SEGMENT: LEGAL REPRESENTATIVE: APPLICATION

HAMILTON, BROOK, SMITH & REYNOLDS, P.C., 530 VIRGINIA ROAD, P.O. BOX 9133, CONCORD, MA, 01742-9133, US

NUMBER OF CLAIMS: 44

EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 2 Drawing Page(s)

LINE COUNT: 2506

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Methods and medical devices for treating a proliferative disorder in a subject, e.g., restenosis in a blood vessel that has been implanted with a stent, employ a bis(thio-hydrazide amide) represented by Structural Formula I or a pharmaceutically acceptable salt or solvate thereof. ##STR1## Y is a covalent bond or an optionally substituted straight chained hydrocarbyl group, or, Y, taken together with both >C=Z groups to which it is bonded, is an optionally substituted aromatic group.

R.sub.1-R.sub.4 are independently --H, an optionally substituted aliphatic group, an optionally substituted aryl group, or R.sub.1 and R.sub.3 taken together with the carbon and nitrogen atoms to which they

are bonded, and/or R.sub.2 and R.sub.4 taken together with the carbon and nitrogen atoms to which they are bonded, form a non-aromatic heterocyclic ring optionally fused to an aromatic ring.

R.sub.7-R.sub.8 are independently --H, an optionally substituted aliphatic group, or an optionally substituted aryl group. Z is O or S.

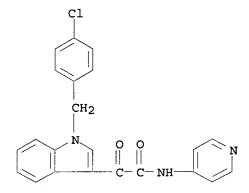
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

204205-90-3, Nascapine

(bis(thiohydrazide amides) for treatment of hyperplasia)

204205-90-3 USPATFULL RN

1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-CN pyridinyl- (CA INDEX NAME)



ANSWER 8 OF 15 USPATFULL on STN

ACCESSION NUMBER:

2006:130825 USPATFULL

TITLE:

Nanoparticulate compositions of tubulin inhibitor

compounds

INVENTOR (S):

Papadopoulos, Pavlos, Antioch, IL, UNITED STATES

Raab, Gerhard, Ronneburg, GERMANY, FEDERAL REPUBLIC OF Doty, Mark J., Grayslake, IL, UNITED STATES Kipp, James E., Wauconda, IL, UNITED STATES

Roessler, Berthold, Halle/Westfalen, GERMANY, FEDERAL

REPUBLIC OF

| | NUMBER | KIND | DATE | |
|---------------------|----------------|------|----------|------|
| | | | | |
| PATENT INFORMATION: | US 2006110462 | A1 | 20060525 | |
| APPLICATION INFO.: | US 2005-266518 | A1 | 20051103 | (11) |

NUMBER DATE 20041108 (60) PRIORITY INFORMATION: US 2004-626036P US 2005-642878P 20050111 (60)

Utility DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

Baxter Healthcare Corporation, One Baxter Parkway -LEGAL REPRESENTATIVE:

DF3-2E, Deerfield, IL, 60015, US

NUMBER OF CLAIMS: 78

EXEMPLARY CLAIM: 1

8 Drawing Page(s)

LINE COUNT:

NUMBER OF DRAWINGS:

2388

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention is directed to novel pharmaceutical compositions comprising nano- and micro-particulate formulations of poorly water soluble tubulin inhibitors of the indole chemical class, preferably N-substituted indol-3-glyoxyamides, and more preferably

N-(Pyridin-4-yl)-[1-(4-chlorobenzyl)-indol-3-yl]glyoxylic acid amide (D-24851), also known as "Indibulin," and methods of making and using such compositions for the treatment of anti-tumor agent resistant cancers and other diseases.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 204205-90-3, D 24851

(Indibulin; particulate compns. of tubulin inhibitors for treatment of resistant cancers and other diseases)

RN 204205-90-3 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

IT 204205-78-7 204205-80-1 204205-81-2 204205-82-3 204205-85-6 204205-86-7

204205-87-8 204205-91-4 204205-92-5

204205-93-6 204205-95-8 204205-96-9

204205-97-0 204205-98-1 204206-01-9

204206-02-0 204206-03-1

(particulate compns. of tubulin inhibitors for treatment of resistant cancers and other diseases)

RN 204205-78-7 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 204205-80-1 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]- α -oxo-N-3-pyridinyl- (CA INDEX NAME)

RN 204205-82-3 USPATFULL CN 1H-Indole-3-acetamide, 1-[(2-chlorophenyl)methyl]- α -oxo-N-3-pyridinyl- (CA INDEX NAME)

RN 204205-85-6 USPATFULL CN 1H-Indole-3-acetamide, N-(2-chloro-3-pyridinyl)-1-[(4-fluorophenyl)methyl]- α -oxo- (CA INDEX NAME)

RN 204205-87-8 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-N-4-pyridinyl-1-(3-pyridinylmethyl)- (CA INDEX NAME)

RN 204205-91-4 USPATFULL CN 1H-Indole-3-acetamide, 1-[(2-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 204205-92-5 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]- α -oxo-N-2-pyridinyl- (CA INDEX NAME)

RN 204205-93-6 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-N-4-pyridinyl-1-(2-pyridinylmethyl)- (CA INDEX NAME)

RN 204205-95-8 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-1-(phenylmethyl)-N-2-pyridinyl- (CA INDEX NAME)

RN 204205-96-9 USPATFULL CN Carbamic acid, [1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-6-yl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 204205-97-0 USPATFULL

CN Carbamic acid, [1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-5-yl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 204205-98-1 USPATFULL

CN Carbamic acid, [1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-6-yl]-, cyclopentyl ester (9CI) (CA INDEX NAME)

RN 204206-01-9 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-5-methoxy- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

$$\begin{array}{c|c} F \\ \hline \\ CH_2 \\ \hline \\ N \\ C-C-NH \\ \hline \end{array}$$

204206-02-0 USPATFULL RN

1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-5-hydroxy- α -oxo-N-CN 4-pyridinyl- (CA INDEX NAME)

204206-03-1 USPATFULL RN

Carbamic acid, [[1-[(4-fluorophenyl)methyl]-3-[oxo(4-CNpyridinylamino)acetyl]-1H-indol-5-yl]methyl]-, ethyl ester (9CI) INDEX NAME)

USPATFULL on STN ANSWER 9 OF 15

2006:4585 USPATFULL ACCESSION NUMBER:

3-glyoxylamideindoles for treating cancer TITLE: Koya, Keizo, Brookline, MA, UNITED STATES INVENTOR(S): Sun, Lijun, Harvard, MA, UNITED STATES

Ono, Mitsunori, Lexington, MA, UNITED STATES

Liang, Guiqing, Concord, MA, UNITED STATES James, David, Cambridge, MA, UNITED STATES

Li, Hao, Brookline, MA, UNITED STATES Xia, Zhi-Qiang, Dedham, MA, UNITED STATES

PATENT ASSIGNEE(S): Synta Pharmaceuticals Corp., Lexington, MA, UNITED

STATES (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: APPLICATION INFO.:

US 2006004044 A1 20060105 US 2005-136074 A1 20050524

RELATED APPLN. INFO.:

US 2005-136074 Al 20050524 (11) Continuation of Ser. No. US 2002-232394, filed on 29

Aug 2002, GRANTED, Pat. No. US 6958348

NUMBER DATE

PRIORITY INFORMATION:

US 2001-322022P 20010913 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HAMILTON, BROOK, SMITH & REYNOLDS, P.C., 530 VIRGINIA

ROAD, P.O. BOX 9133, CONCORD, MA, 01742-9133, US

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

26 1-20

NUMBER OF DRAWINGS:

2 Drawing Page(s)

LINE COUNT:

1010

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Disclosed is an anti-cancer compound represented by Structural Formula (I): ##STR1## The variables in Structural Formula (I) are described hereinbelow. Also disclosed is a pharmaceutical composition comprising a pharmaceutically acceptable carrier or diluent and a compound represented by Structural Formula (I) (preferably an effective amount). Also disclosed is a method of treating a subject with cancer by administering to the subject an effective amount of a compound represented by Structural Formula (I).

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 501921-60-4P 501921-65-9P

(preparation of glyoxlylamide indoles as anticancer agents useful against multidrug-resistant cancer cells)

RN 501921-60-4 USPATFULL

CN 1H-Indole-3-acetamide, N-[5-(aminocarbonyl)-2-pyridinyl]-1-[(4-chlorophenyl)methyl]-α-οxο- (CA INDEX NAME)

RN

L6 ANSWER 10 OF 15 USPATFULL on STN

ACCESSION NUMBER:

2005:268778 USPATFULL

TITLE:

Novel compounds and methods of use thereof

INVENTOR(S):

Chen, Chiung-Tong, Taipei, TAIWAN, PROVINCE OF CHINA Chen, Shu-Jen, Taipei, TAIWAN, PROVINCE OF CHINA Hsu, Ming-Chu, Taipei, TAIWAN, PROVINCE OF CHINA Hwang, Der-Ren, Taipei, TAIWAN, PROVINCE OF CHINA Li, Wen-Tai, Taipei, TAIWAN, PROVINCE OF CHINA Lin, Chu-Chung, Taipei, TAIWAN, PROVINCE OF CHINA

KIND NUMBER DATE

PATENT INFORMATION:

US 2005234098

A1 20051020

APPLICATION INFO.:

US 2005-145628

A1 20050606 (11)

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2002-310711, filed on 5 Dec

2002, GRANTED, Pat. No. US 6903104

NUMBER DATE

PRIORITY INFORMATION:

US 2001-337962P

20011206 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

FISH & RICHARDSON PC, P.O. BOX 1022, MINNEAPOLIS, MN,

55440-1022, US

NUMBER OF CLAIMS:

30

EXEMPLARY CLAIM: NUMBER OF DRAWINGS: 1-37 2 Drawing Page(s)

LINE COUNT:

2031

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention relates to novel heteroatom containing compounds and compositions thereof, and their use for the prevention and treatment of disease. The invention also provides for methods of making the compounds. The invention is based on the discovery that certain heteroatom containing compounds, 3-oxoacetamideindolyl compounds, have potent anticancer, cytotoxic, and anti-angiogenic activity.

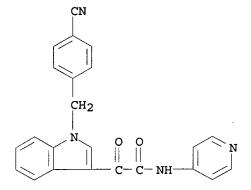
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 501921-65-9P, N-(4-Pyridyl)-2-[1-(4-cyanobenzyl)-1H-indol-3-yl]-2oxoacetamide

(preparation of (3-indolyl)oxoacetamide derivs. as angiogenesis inhibitors and anticancer agents)

501921-65-9 USPATFULL RN

1H-Indole-3-acetamide, 1-[(4-cyanophenyl)methyl]- α -oxo-N-4-pyridinyl-CN (CA INDEX NAME)



L6 ANSWER 11 OF 15 USPATFULL on STN

ACCESSION NUMBER: 2005:171786 USPATFULL

TITLE: IAP nucleobase oligomers and oligomeric complexes and

uses thereof

INVENTOR(S): LaCasse, Eric, Ottawa, CANADA

McManus, Daniel, Ottawa, CANADA

APPLICATION INFO.: US 2004-975974 A1 20041028 (10)

NUMBER DATE

PRIORITY INFORMATION: US 2003-516192P 20031030 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: CLARK & ELBING LLP, 101 FEDERAL STREET, BOSTON, MA,

02110, US

NUMBER OF CLAIMS: 48 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 15 Drawing Page(s)

LINE COUNT: 3022

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides nucleobase oligomers and oligomer complexes that inhibit expression of an IAP polypeptide, and methods for using them to induce apoptosis in a cell. The nucleobase oligomers and oligomer complexes of the present invention may also be used to form pharmaceutical compositions. The invention also features methods for enhancing apoptosis in a cell by administering a nucleobase oligomer or oligomer complex of the invention in combination with a chemotherapeutic or chemosensitizing agent.

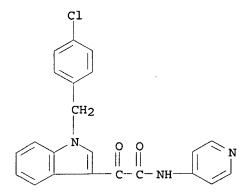
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 204205-90-3, D 24851

(human protein IAP (inhibitor of apoptosis protein) nucleobase oligomers, including dsRNA, shRNA, and siRNA, and their use for enhancing apoptosis in cancer therapy)

RN 204205-90-3 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)



ANSWER 12 OF 15 USPATFULL on STN

ACCESSION NUMBER:

2004:221896 USPATFULL

TITLE:

Indolyl-3-glyoxylic acid derivatives having

therapeutically valuable properties

INVENTOR (S):

Nickel, Bernd, Muhltal, GERMANY, FEDERAL REPUBLIC OF

Bacher, Gerald, Heidelberg, GERMANY, FEDERAL REPUBLIC

Klenner, Thomas, Ingelheim, GERMANY, FEDERAL REPUBLIC

Beckers, Thomas, Frankfurt, GERMANY, FEDERAL REPUBLIC

Emig, Peter, Bruchkobel, GERMANY, FEDERAL REPUBLIC OF Engel, Jurgen, Alzenau, GERMANY, FEDERAL REPUBLIC OF

Bruyneel, Erik, Harelbeke, BELGIUM

Kamp, Gunter, Munster, GERMANY, FEDERAL REPUBLIC OF Peters, Kirsten, Munster, GERMANY, FEDERAL REPUBLIC OF

PATENT ASSIGNEE(S):

Baxter Healthcare SA, Wallisellen, SWITZERLAND

(non-U.S. corporation)

| | NUMBER | KIND | DATE |
|----|------------|------|----------|
| | | | - |
| US | 2004171668 | A1 | 20040902 |
| | | | |

PATENT INFORMATION: APPLICATION INFO.:

US 2003-686809 A1 20031017 (10)

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2000-492531, filed on 27

Jan 2000, GRANTED, Pat. No. US 6693119

Continuation-in-part of Ser. No. US 1999-285058, filed

on 2 Apr 1999, GRANTED, Pat. No. US 6232327

NUMBER DATE **----**PRIORITY INFORMATION: DE 1999-19946301 19990828 DE 1998-19814838 19980402

DOCUMENT TYPE: FILE SEGMENT:

Utility

APPLICATION

LEGAL REPRESENTATIVE:

PILLSBURY WINTHROP, LLP, P.O. BOX 10500, MCLEAN, VA,

22102

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

13 1

9 Drawing Page(s)

NUMBER OF DRAWINGS: LINE COUNT:

570

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to the use of N-substituted indole-3-glyoxylamides of the general Formula I: ##STR1##

and to pharmaceutical compositions having antitumor action.

CAS INDEXING IS AVAILABLE FOR THIS PATENT. IT 204205-78-7P 204205-79-8P 204205-80-1P

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204205-81-2P 204205-82-3P 204205-85-6P
      204205-86-7P 204205-87-8P 204205-90-3P
      204205-91-4P 204205-92-5P 204205-93-6P
      204205-95-8P 204205-96-9P 204205-97-0P
      204206-01-9P 204206-03-1P 245661-24-9P
      245661-25-0P 245661-26-1P 245661-28-3P
      245661-29-4P 245661-30-7P 245661-31-8P
      245661-38-5P 245661-39-6P 245661-41-0P
      245661-42-1P 245661-43-2P 245661-47-6P
      245661-48-7P 245661-49-8P 245661-50-1P
      245661-51-2P 245661-52-3P 245661-53-4P
      245661-54-5P 245661-55-6P
        (preparation of indolylglyoxylamides as antitumor agents)
RN
     204205-78-7 USPATFULL
CN
     1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-\alpha-oxo-N-4-
       pyridinyl- (CA INDEX NAME)
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RN 204205-79-8 USPATFULL CN 1H-Indole-3-acetamide, 1-methyl- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 204205-80-1 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]- α -oxo-N-3-pyridinyl- (CA INDEX NAME)

RN 204205-81-2 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-1-(phenylmethyl)-N-3-pyridinyl- (CA INDEX NAME)

RN 204205-82-3 USPATFULL CN 1H-Indole-3-acetamide, 1-[(2-chlorophenyl)methyl]- α -oxo-N-3-pyridinyl- (CA INDEX NAME)

RN 204205-85-6 USPATFULL
CN 1H-Indole-3-acetamide, N-(2-chloro-3-pyridinyl)-1-[(4-fluorophenyl)methyl]α-οxo- (CA INDEX NAME)

RN 204205-87-8 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-N-4-pyridinyl-1-(3-pyridinylmethyl)- (CA INDEX NAME)

RN 204205-90-3 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 204205-91-4 USPATFULL CN 1H-Indole-3-acetamide, 1-[(2-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 204205-92-5 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]- α -oxo-N-2-pyridinyl- (CA INDEX NAME)

RN 204205-93-6 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-N-4-pyridinyl-1-(2-pyridinylmethyl)- (CA INDEX NAME)

RN 204205-96-9 USPATFULL

CN Carbamic acid, [1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-6-yl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 204205-97-0 USPATFULL

CN Carbamic acid, [1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-5-yl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 204206-01-9 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-5-methoxy- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 204206-03-1 USPATFULL

CN Carbamic acid, [[1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-5-yl]methyl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 245661-24-9 USPATFULL

CN 1H-Indole-3-acetamide, 5-fluoro-1-[(4-fluorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-25-0 USPATFULL CN 1H-Indole-3-acetamide, 1-[(2-bromophenyl)methyl]- α -oxo-N-4-pyridinyl-(CA INDEX NAME)

RN 245661-26-1 USPATFULL CN 1H-Indole-3-acetamide, 1-[(3-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-28-3 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-bromophenyl)methyl]- α -oxo-N-4-pyridinyl-(CA INDEX NAME)

RN 245661-29-4 USPATFULL CN 1H-Indole-3-acetamide, 1-[(3-fluorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-30-7 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]-N-(1-oxido-4-pyridinyl)- α -oxo- (CA INDEX NAME)

RN 245661-31-8 USPATFULL CN 1H-Indole-3-acetamide, N-(4-amino-3-pyridinyl)-1-[(4-fluorophenyl)methyl]- α -oxo- (CA INDEX NAME)

RN 245661-38-5 USPATFULL
CN Carbamic acid, [[1-[(4-fluorophenyl)methyl]-3-[oxo(4pyridinylamino)acetyl]-1H-indol-5-yl]methyl]-, methyl ester (9CI) (CA
INDEX NAME)

RN 245661-39-6 USPATFULL
CN Carbamic acid, [[1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-5-yl]methyl]-, 2-methylpropyl ester (9CI) (CA INDEX NAME)

RN 245661-41-0 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-6-nitro- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-42-1 USPATFULL

CN lH-Indole-3-acetamide, l-[(4-fluorophenyl)methyl]-5-nitro- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & \\ & & \\ &$$

RN 245661-43-2 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(2-fluorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-47-6 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-α-oxo-N-4-pyridinyl-, monohydrochloride (9CI) (CA INDEX NAME)

● HCl

RN 245661-48-7 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 204205-90-3 CMF C22 H16 Cl N3 O2

CM 2

CRN 76-05-1 CMF C2 H F3 O2

RN 245661-49-8 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-methoxyphenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-50-1 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(2,6-dichlorophenyl)methyl]-α-oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-51-2 USPATFULL

CN Carbamic acid, [1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-5-yl]-, 2-methylpropyl ester (9CI) (CA INDEX NAME)

RN 245661-52-3 USPATFULL

CN lH-Indole-3-acetamide, α -oxo-N-4-pyridinyl-1-[[4-(trifluoromethyl)phenyl]methyl]- (CA INDEX NAME)

RN 245661-53-4 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]-N-(3,5-dichloro-2-pyridinyl)-α-oxo- (CA INDEX NAME)

RN 245661-54-5 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-methylphenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-55-6 USPATFULL

CN lH-Indole-3-acetamide, l-[(2,4-dichlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

L6 ANSWER 13 OF 15 USPATFULL on STN

ACCESSION NUMBER:

2003:258429 USPATFULL

TITLE:

Novel compounds and methods of use thereof

INVENTOR (S):

Chen, Chiung-Tong, Taipei, TAIWAN, PROVINCE OF CHINA Chen, Shu-Jen, Taipei, TAIWAN, PROVINCE OF CHINA Hsu, Ming-Chu, Taipei, TAIWAN, PROVINCE OF CHINA Hwang, Der-Ren, Taipei, TAIWAN, PROVINCE OF CHINA Li, Wen-Tai, Taipei, TAIWAN, PROVINCE OF CHINA Lin, Chu-Chung, Taipei, TAIWAN, PROVINCE OF CHINA

| | NUMBER | KIND | DATE | |
|---------------------|-----------------------------|----------|----------|------|
| PATENT INFORMATION: | US 2003181482 US 6903104 | A1 B2 | 20030925 | |
| APPLICATION INFO.: | US 2002-310711 | A1 | 20021205 | (10) |

NUMBER DATE

PRIORITY INFORMATION:

US 2001-337962P 20011206 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

JEFFREY D. HSI, Fish & Richarson P.C., 225 Franklin

Street, Boston, MA, 02110-2804

NUMBER OF CLAIMS:

TYPINDIADY OF THE

37 1

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

2 Drawing Page(s)

LINE COUNT:

2068

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention relates to novel heteroatom containing compounds and compositions thereof, and their use for the prevention and treatment of disease. The invention also provides for methods of making the compounds. The invention is based on the discovery that certain heteroatom containing compounds, 3-oxoacetamideindolyl compounds, have potent anticancer, cytotoxic, and anti-angiogenic activity.

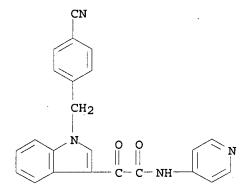
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 501921-65-9P, N-(4-Pyridyl)-2-[1-(4-cyanobenzyl)-1H-indol-3-yl]-2-oxoacetamide

(preparation of (3-indolyl)oxoacetamide derivs. as angiogenesis inhibitors and anticancer agents)

RN 501921-65-9 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-cyanophenyl)methyl]- α -oxo-N-4-pyridinyl-(CA INDEX NAME)



ANSWER 14 OF 15 USPATFULL on STN

ACCESSION NUMBER:

2003:166653 USPATFULL

TITLE:

Indoly1-3-glyoxylic acid derivatives having

therapeutically valuable properties

INVENTOR(S):

Nickel, Bernd, Muhltal, GERMANY, FEDERAL REPUBLIC OF

Bacher, Gerald, Heidelberg, GERMANY, FEDERAL REPUBLIC

Klenner, Thomas, Ingelheim, GERMANY, FEDERAL REPUBLIC

OF

Beckers, Thomas, Frankfurt, GERMANY, FEDERAL REPUBLIC

OF

Emig, Peter, Bruchkobel, GERMANY, FEDERAL REPUBLIC OF Engel, Jurgen, Alzenau, GERMANY, FEDERAL REPUBLIC OF

Bruyneel, Erik, Harelbeke, BELGIUM

Kamp, Gunter, Munster, GERMANY, FEDERAL REPUBLIC OF Peters, Kirsten, Munster, GERMANY, FEDERAL REPUBLIC OF

| | NUMBER | KIND | DATE | |
|---------------------|----------------|------|----------|-----|
| PATENT INFORMATION: | US 2003114511 | A1 | 20030619 | |
| PAIENT INFORMATION: | US 6693119 | B2 | 20030619 | |
| APPLICATION INFO.: | US 2000-492531 | Al | 20000127 | (9) |

NUMBER DATE

PRIORITY INFORMATION:

DE 1998-19814838 19980402 DE 1999-19946301 19990928

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

PILLSBURY WINTHROP, LLP, P.O. BOX 10500, MCLEAN, VA,

22102

NUMBER OF CLAIMS:

13

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

1

9 Drawing Page(s)

LINE COUNT:

576

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The object of the invention is then to widen the field of use of AB N-substituted indole-3-glyoxylamides and thus to enrich the available pharmaceutical wealth. The possibility of a lower, longer-lasting and better-tolerable medication for the class of substances having antitumor action described in German Patent Application 19814 838.0 should thus be opened up. In particular, the disadvantageous development of resistance, as is known of many antitumor agents, should be circumvented.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 204205-78-7P 204205-79-8P 204205-80-1P

204205-81-2P 204205-82-3P 204205-85-6P

204205-86-7P 204205-87-8P 204205-90-3P

204205-91-4P 204205-92-5P 204205-93-6P 204205-95-8P 204205-96-9P 204205-97-0P 204206-01-9P 204206-03-1P 245661-24-9P 245661-25-0P 245661-26-1P 245661-28-3P 245661-29-4P 245661-30-7P 245661-31-8P 245661-38-5P 245661-39-6P 245661-41-0P 245661-42-1P 245661-43-2P 245661-47-6P 245661-48-7P 245661-49-8P 245661-50-1P 245661-51-2P 245661-52-3P 245661-53-4P 245661-54-5P 245661-55-6P (preparation of indolylglyoxylamides as antitumor agents) 204205-78-7 USPATFULL RN1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]- α -oxo-N-4-CNpyridinyl- (CA INDEX NAME)

RN 204205-79-8 USPATFULL CN 1H-Indole-3-acetamide, 1-methyl- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 204205-80-1 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]- α -oxo-N-3-pyridinyl- (CA INDEX NAME)

RN 204205-81-2 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-1-(phenylmethyl)-N-3-pyridinyl- (CA INDEX NAME)

RN 204205-85-6 USPATFULL CN 1H-Indole-3-acetamide, N-(2-chloro-3-pyridinyl)-1-[(4-fluorophenyl)methyl]- α -oxo- (CA INDEX NAME)

RN 204205-86-7 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-1-(phenylmethyl)-N-4-pyridinyl- (CA INDEX NAME)

RN 204205-92-5 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]- α -oxo-N-2-pyridinyl- (CA INDEX NAME)

RN 204205-93-6 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-N-4-pyridinyl-1-(2-pyridinylmethyl)- (CA INDEX NAME)

RN 204205-95-8 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-1-(phenylmethyl)-N-2-pyridinyl- (CA INDEX NAME)

RN 204205-97-0 USPATFULL

CN Carbamic acid, [1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-5-yl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 204206-01-9 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-5-methoxy-α-οxο-N-4-pyridinyl- (CA INDEX NAME)

RN 204206-03-1 USPATFULL

CN Carbamic acid, [[1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-5-yl]methyl]-, ethyl ester (9CI) (CAINDEX NAME)

245661-24-9 USPATFULL RN

 $1 \\ \\ \text{H-Indole-3-acetamide, 5-fluoro-1-[(4-fluorophenyl)methyl]-} \\ \\ \alpha - oxo-\\ \\ \text{N-4-parameters}$ CN pyridinyl- (CA INDEX NAME)

RN 245661-25-0 USPATFULL

1H-Indole-3-acetamide, 1-[(2-bromophenyl)methyl]- α -oxo-N-4-pyridinyl-CN(CA INDEX NAME)

245661-26-1 USPATFULL RN

1H-Indole-3-acetamide, 1-[(3-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME) CN

RN 245661-28-3 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-bromophenyl)methyl]- α -oxo-N-4-pyridinyl-(CA INDEX NAME)

RN 245661-29-4 USPATFULL CN 1H-Indole-3-acetamide, 1-[(3-fluorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-30-7 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]-N-(1-oxido-4-pyridinyl)- α -oxo- (CA INDEX NAME)

RN 245661-31-8 USPATFULL CN 1H-Indole-3-acetamide, N-(4-amino-3-pyridinyl)-1-[(4-fluorophenyl)methyl]- α -oxo- (CA INDEX NAME)

RN 245661-41-0 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-6-nitro-α-oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-42-1 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-5-nitro-α-oxo-N-4-pyridinyl- (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & \\ & \\ & \\ & \\ & \\ & \\ & \\ & \\ & \\ & \\ & \\ & \\ & \\ \end{array}$$

RN 245661-43-2 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(2-fluorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-47-6 USPATFULL
CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-α-oxo-N-4pyridinyl-, monohydrochloride (9CI) (CA INDEX NAME)

● HCl

RN 245661-48-7 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 204205-90-3 CMF C22 H16 Cl N3 O2 .

CRN 76-05-1 CMF C2 H F3 O2

RN 245661-49-8 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-methoxyphenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-50-1 USPATFULL CN 1H-Indole-3-acetamide, 1-[(2,6-dichlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 245661-51-2 USPATFULL
CN Carbamic acid, [1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]1H-indol-5-yl]-, 2-methylpropyl ester (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ \hline \\ i-BuO-C-NH & & & \\ \end{array}$$

245661-52-3 USPATFULL RN CN

1H-Indole-3-acetamide, α -oxo-N-4-pyridinyl-1-[[4-(trifluoromethyl)phenyl]methyl] - (CA INDEX NAME)

RN 245661-53-4 USPATFULL

1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]-N-(3,5-dichloro-2-CNpyridinyl) $-\alpha$ -oxo- (CA INDEX NAME)

RN245661-54-5 USPATFULL

1H-Indole-3-acetamide, 1-[(4-methylphenyl)methyl]- α -oxo-N-4-CNpyridinyl- (CA INDEX NAME)

245661-55-6 USPATFULL RN

1H-Indole-3-acetamide, 1-[(2,4-dichlorophenyl)methyl]- α -oxo-N-4-CNpyridinyl- (CA INDEX NAME)

ANSWER 15 OF 15 USPATFULL on STN L6

ACCESSION NUMBER: 2003:134662 USPATFULL

TITLE:

INVENTOR(S):

3-glyoxlylamideindoles for treating cancer Koya, Keizo, Brookline, MA, UNITED STATES

Sun, Lijun, Harvard, MA, UNITED STATES

Ono, Mitsunori, Lexington, MA, UNITED STATES Liang, Guiqing, Concord, MA, UNITED STATES James, David, Cambridge, MA, UNITED STATES Li, Hao, Brookline, MA, UNITED STATES

Xia, Zhi-Qiang, Dedham, MA, UNITED STATES SBR Pharmaceuticals Corp., Lexington, MA (U.S.

corporation)

| | NUMBER | KIND | DATE | |
|---------------------|-----------------------------|----------|----------|------|
| PATENT INFORMATION: | US 2003092751 US 6958348 | A1 B2 | 20030515 | |
| APPLICATION INFO.: | US 2002-232394 | A1 | 20020829 | (10) |

NUMBER DATE

PRIORITY INFORMATION:

PATENT ASSIGNEE(S):

US 2001-322022P 20010913 (60)

DOCUMENT TYPE:

Utility APPLICATION

FILE SEGMENT: LEGAL REPRESENTATIVE:

HAMILTON, BROOK, SMITH & REYNOLDS, P.C., 530 VIRGINIA

ROAD, P.O. BOX 9133, CONCORD, MA, 01742-9133

NUMBER OF CLAIMS:

42

EXEMPLARY CLAIM:

1

NUMBER OF DRAWINGS: 2 Drawing Page(s)

LINE COUNT: 1151

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Disclosed is an anti-cancer compound represented by Structural Formula (I): ##STR1##

The variables in Structural Formula (I) are described hereinbelow. Also disclosed is a pharmaceutical composition comprising a pharmaceutically acceptable carrier or diluent and a compound represented by Structural Formula (I) (preferably an effective amount). Also disclosed is a method of treating a subject with cancer by administering to the subject an effective amount of a compound represented by Structural Formula (I).

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 501921-60-4P 501921-65-9P

(preparation of glyoxlylamide indoles as anticancer agents useful against multidrug-resistant cancer cells)

RN 501921-60-4 USPATFULL

CN 1H-Indole-3-acetamide, N-[5-(aminocarbonyl)-2-pyridinyl]-1-[(4-chlorophenyl)methyl]- α -oxo- (CA INDEX NAME)

=> d his (FILE 'HOME' ENTERED AT 15:23:09 ON 24 JAN 2008) FILE 'REGISTRY' ENTERED AT 15:23:20 ON 24 JAN 2008 STRUCTURE UPLOADED L1 L2 143 S L1 FULL FILE 'MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 15:23:58 ON 24 JAN 2008 185 S L2 L358 S L3 AND (CANCER? OR TUMOR?) L413 S L4 AND ANGIOGENESIS L5 15 S L4 AND (MULTIDRUG OR MULTI-DRUG) L6 => s 14 not py>2000 1 L4 NOT PY>2000 L7 => d 17 ibib, abs, hitstr ANSWER 1 OF 1 CAPLUS COPYRIGHT 2008 ACS on STN 2000:814353 CAPLUS ACCESSION NUMBER: DOCUMENT NUMBER: 133:359224 Fatty acid-N-substituted indol-3-glyoxylamide TITLE: compositions as antitumor agents Bradley, Matthews O.; Swindell, Charles S.; Anthony, Forrest; Webb, Nigel L.; Fisher, Mark INVENTOR(S): Protarga, Inc., USA PCT Int. Appl., 48 pp. PATENT ASSIGNEE(S): SOURCE: CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: DATE KTND APPLICATION NO. האתבאת אה אדעם

| | PATENT NO. | | | | | KIND DATE | | | APPLICATION NO. | | | | | | DATE | | | |
|---------------|---------------------------------------|---------|-------|-----|------------|-----------|-----|------|-----------------|-----|------|-------|-------|-----------|------|-----|------|-----|
| | | | | | | | - | | - | | | | | · | | - | | |
| | WO | 2000 | 0678 | 02 | | A1 | | 2000 | 1116 | 1 | WO 2 | 000-1 | US12' | 752 | | 2 | 0000 | 510 |
| | | W: | ΑE, | AG, | AL, | AM, | ΑT, | AU, | ΑZ, | BA, | BB, | BG, | BR, | BY, | CA, | CH, | CN, | CR, |
| | | | | | | | | DZ, | | | | | | | | | | |
| | | | ID, | IL, | IN, | IS, | JΡ, | KE, | KG, | KP, | KR, | KZ, | LC, | LK, | LR, | LS, | LT, | LU, |
| | | | | | | | | MN, | | | | | | | | | | |
| | | | SG, | SI, | SK, | SL, | TJ, | TM, | TR, | TT, | TZ, | UA, | ÜĠ, | UZ, | VN, | YU, | ZA, | ZW, |
| | | | AM, | ΑZ, | BY, | KG, | ΚZ, | MD, | RU, | TJ, | TM | | | | | | | |
| | | RW: | GH, | GM, | KE, | LS, | MW, | SD, | SL, | SZ, | TZ, | ŪG, | ZW, | ΑT, | BE, | CH, | CY, | DE, |
| | | | DK, | ES, | FI, | FR, | GB, | GR, | ΙE, | IT, | LU, | MC, | NL, | PT, | SE, | BF, | ΒJ, | CF, |
| | | | CG, | CI, | CM, | GΑ, | GN, | GW, | ML, | MR, | ΝE, | SN, | TD, | TG | | | | |
| AU 2000048342 | | | | | A 20001121 | | | 1121 | AU 2000-48342 | | | | 2 | 20000510 | | | | |
| PRIO | RIORITY APPLN. INFO.: US 1999-133292P | | | | | ; | P 1 | 9990 | 510 | | | | | | | | | |
| | | | | | | | | | | 1 | WO 2 | 000-1 | US12' | 752 | | ₩ 2 | 0000 | 510 |
| OTHE | 2 50 | TIRCE | (S) · | | | MAR | PAT | 133: | 3592 | 2.4 | | | | | | | | |

OTHER SOURCE(S): MARPAT 133:359224

GI

The present invention pertains to N-substituted indol-3-glyoxylamides that are conjugates of fatty acids and conjugates of I. The conjugates are useful in treating cancer. In an example taxoprexin completely eliminated all measureable tumors in 7 out of 8 mice at 120 mg/kg/day for 5 days while paclitaxel retarded tumor growth for about 4 days.

IT 204205-90-3D, conjugates, with antitumor agents RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(fatty acid-N-substituted indol-3-glyoxylamide compns. as antitumor agents)

RN 204205-90-3 CAPLUS

CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> d his

(FILE 'HOME' ENTERED AT 15:23:09 ON 24 JAN 2008)

7

FILE 'REGISTRY' ENTERED AT 15:23:20 ON 24 JAN 2008

L1 STRUCTURE UPLOADED

L2 143 S L1 FULL

FILE 'MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 15:23:58 ON 24 JAN 2008

L3 185 S L2

58 S L3 AND (CANCER? OR TUMOR?) L4L5 13 S L4 AND ANGIOGENESIS 15 S L4 AND (MULTIDRUG OR MULTI-DRUG) L6 1 S L4 NOT PY>2000 1.7 => s 13 not py>2000 14 L3 NOT PY>2000 L8 => d 18 1-14 ibib, abs, hitstr ANSWER 1 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN 2001:30560 CAPLUS ACCESSION NUMBER: 134:221365 DOCUMENT NUMBER: The effect of selective and non-selective TITLE: phosphodiesterase inhibitors on allergen- and leukotriene C4-induced contractions in passively sensitized human airways Schmidt, Dunja T.; Watson, Nikki; Dent, Gordon; AUTHOR(S): Ruhlmann, Elke; Branscheid, Detlev; Magnussen, Helgo; Rabe, Klaus F. Department of Pulmonology, Leiden University Medical CORPORATE SOURCE: Centre, Leiden, NL-2333 ZA, Neth. British Journal of Pharmacology (2000), 131(8), SOURCE: 1607-1618 CODEN: BJPCBM; ISSN: 0007-1188 Nature Publishing Group PUBLISHER: DOCUMENT TYPE: Journal English LANGUAGE: Non-selective inhibitors of cyclic nucleotide phosphodiesterase (PDE) block allergen-induced contraction of passively sensitized human airways in vitro by a dual mechanism involving a direct relaxant effect on smooth muscle and inhibition of histamine and cysteinyl leukotriene (LT) release from airways. We investigated the effects of non-selective PDE inhibitors and selective inhibitors of PDE3 and PDE4 in order to determine the involvement of PDE isoenzymes in the suppression of allergic bronchoconstriction. Macroscopically normal airways from 76 patients were sensitized with IgE-rich sera (>250 u ml-1) containing specific antibodies against allergen (Dermatophagoides farinae). Contractile responses of bronchial rings were assessed using standard organ bath techniques. Passive sensitization caused increased contractile responses to allergen, histamine and LTC4. Non-selective PDE inhibitors (theophylline, 3-isobutyl-1-methylxanthine [IBMX]), a PDE3-selective inhibitor (motapizone), PDE4-selective inhibitors (RP73401, rolipram, AWD 12-281) and a mixed PDE3/4 inhibitor (zardaverine) all significantly relaxed inherent bronchial tone at resting tension and to a similar degree. Theophylline, IBMX, zardaverine and the combination of motapizone and RP73401 inhibited the contractile responses to allergen and LTC4. Pre-treatment with motapizone, RP73401, rolipram or the methylxanthine adenosine receptor antagonist, 8-phenyltheophylline, did not significantly decrease responses to either allergen or LTC4. We conclude that combined inhibition of PDE3 and PDE4, but not selective inhibition of either isoenzyme or antagonism of adenosine receptors, is effective in suppressing allergen-induced contractions of passively sensitized human airways. The relationship between allergen- and LTC4-induced responses suggests that PDE inhibitors with PDE3 and PDE4 selectivity are likely to act in part through inhibition of mediator release and not simply through direct relaxant actions on airway smooth muscle. IT 257892-33-4, AWD 12-281 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (phosphodiesterase inhibitors in allergen- and leukotriene C4-induced contractions in sensitized human airways)

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-

257892-33-4 CAPLUS

RN

CN

52 THERE ARE 52 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 2 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2000:814353 CAPLUS

DOCUMENT NUMBER: 133:359224

Fatty acid-N-substituted indol-3-glyoxylamide TITLE:

compositions as antitumor agents

Bradley, Matthews O.; Swindell, Charles S.; Anthony, Forrest; Webb, Nigel L.; Fisher, Mark INVENTOR(S):

Protarga, Inc., USA PCT Int. Appl., 48 pp. PATENT ASSIGNEE(S): SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| | PATEN | 1 T | 10. | | | KIN | D | DATE | | | APPL | ICAT | ION I | NO. | | D | ATE | | |
|---------------|--------|------|------|------------|-----|------|--------|---------------|----------------|-----|------|----------|-------|-------|-----|-----|-------|-----|--|
| | | | | | | | - | | - - | | | <i>-</i> | | | | - | | | |
| | WO 20 | 000 | 6780 | 02 | | A1 | | 2000 | 1116 | • | WO 2 | 000- | US12 | 752 | | 2 | 0000 | 510 | |
| | W | ∛: | AE, | AG, | AL, | AM, | AT, | ΑU, | ΑZ, | BA, | BB, | BG, | BR, | BY, | CA, | CH, | CN, | CR, | |
| | | | CU, | CZ, | DE, | DK, | DM, | DZ, | EE, | ES, | FI, | GB, | GD, | GE, | GH, | GM, | HR, | HU, | |
| | | | ID, | IL, | IN, | IS, | JP, | KE, | KG, | KP, | KR, | ΚZ, | LC, | LK, | LR, | LS, | LT, | LU, | |
| | | | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | NO, | NZ, | PL, | PT, | RO, | RU, | SD, | SE, | |
| | | | SG, | SI, | SK, | SL, | ΤĴ, | TM, | TR, | TT, | ΤZ, | UA, | ŪĠ, | UΖ, | VN, | YU, | ZA, | ZW, | |
| | | | AM, | AZ; | BY, | KG, | ΚZ, | MD, | RU, | ТJ, | TM | | | | | | | | |
| | R | : WS | GH, | GM, | KE, | LS, | MW, | SD, | SL, | SZ, | TZ, | UG, | ZW, | ΑT, | BE, | CH, | CY, | DΕ, | |
| | | | DK, | ES, | FI, | FR, | GB, | GR, | ΙE, | IT, | LU, | MC, | NL, | PT, | SE, | BF, | ВJ, | CF, | |
| | | | CG, | CI, | CM, | GA, | GN, | GW, | ML, | MR, | NE, | SN, | TD, | TG | | | | | |
| AU 2000048342 | | | | A 20001121 | | | 1121 | AU 2000-48342 | | | | | 2 | 0000! | 510 | | | | |
| PRIO | RITY A | APPI | .N. | INFO | . : | | | | | • | US 1 | 999- | 1332 | 92P | | P 1 | 9990! | 510 | |
| | | | | | | | | | | 1 | WO 2 | 000-1 | JS12 | 752 | 1 | W 2 | 0000 | 510 | |
| OTHER | COLLE | 000 | (0). | | | MADE | ייייעם | 122. | 2 5 0 2 1 | 2.4 | | | | | | | | | |

OTHER SOURCE(S): MARPAT 133:359224

GI

The present invention pertains to N-substituted indol-3-glyoxylamides that AB are conjugates of fatty acids and conjugates of I. The conjugates are useful in treating cancer. In an example taxoprexin completely eliminated all measureable tumors in 7 out of 8 mice at 120 mg/kg/day for 5 days while paclitaxel retarded tumor growth for about 4 days.

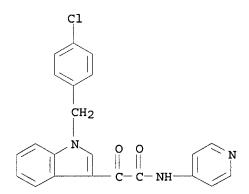
204205-90-3D, conjugates, with antitumor agents RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(fatty acid-N-substituted indol-3-glyoxylamide compns. as antitumor agents)

RN 204205-90-3 CAPLUS

1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-CN pyridinyl- (CA INDEX NAME)

Ι



THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

CAPLUS COPYRIGHT 2008 ACS on STN L8 ANSWER 3 OF 14

2000:55462 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 132:202635

A peptidic binding site model for PDE 4 inhibitors TITLE:

Polymeropoulos, Emmanuel E.; Hofgen, Norbert AUTHOR(S):

CORPORATE SOURCE: Department of Chemical Research, Corporate R and D

ASTA Medica Group, Frankfurt, D-60314, Germany

Quantitative Structure-Activity Relationships (1999), SOURCE:

18(6), 543-547

CODEN: QSARDI; ISSN: 0931-8771

PUBLISHER: Wiley-VCH Verlag GmbH

Journal DOCUMENT TYPE:

LANGUAGE: English The pseudoreceptor modeling program PrGen was used to construct a peptidic binding site model for phosphodiesterase 4 inhibitors. A training set of 21 diverse compds. (rolipram, nitraquazone and xanthine derivs., imidazo pyrido pyrazinones and 5-oxyindoles) was used to construct the binding site surrogate consisting of five amino acid residues, a Zn+2 cofactor and an envelope of charged virtual particles. The model was validated by predicting the free energies of binding ΔGpred0 of ten ligands (rolipram, imidazo pyrido pyrazinones and 5-oxyindoles). In seven cases the prediction was satisfactory. The rms deviation [4] in ΔG0 is 0.16 and 1.82 kcal/mol-resulting in an uncertainty in IC50 (or Ki) of 1.32 and 22.81-for the training and the test set resp., while the corresponding maximal prediction errors in ΔGpred0 were 0.27 kcal/mol and 4.50 kcal/mol.

CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-5-hydroxy-α-oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 247584-23-2 CAPLUS
CN 1H-Indole-3-acetamide, 1-[(2,6-difluorophenyl)methyl]-5-hydroxy-αoxo-N-4-pyridinyl- (CA INDEX NAME)

RN 247584-24-3 CAPLUS
CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(2,6-difluorophenyl)methyl]-5-hydroxy-α-oxo- (CA INDEX NAME)

RN 247584-27-6 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-5-hydroxy-1-(1-methylethyl)- α -oxo- (CA INDEX NAME)

RN 247584-34-5 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-methoxy- α -oxo- (CA INDEX NAME)

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-οxο- (CA INDEX NAME)

RN 260265-54-1 CAPLUS

CN 1H-Indole-3-acetamide, N-(2,6-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-οxο- (CA INDEX NAME)

RN 260265-55-2 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-5-methoxy-1-(1-methylethyl)- α -oxo- (CA INDEX NAME)

RN 260265-56-3 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-5-methoxy-1-[(3-nitrophenyl)methyl]-α-οxο- (CA INDEX NAME)

RN 260265-57-4 CAPLUS CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-5-hydroxy-1-[(3-nitrophenyl)methyl]- α -oxo- (CA INDEX NAME)

$$\begin{array}{c|c} O_2N \\ \hline \\ CH_2 \\ \hline \\ N \\ O \\ C-C-NH \\ \hline \\ C1 \\ \end{array}$$

RN (260265-58-5 CAPLUS
CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(2,6-difluorophenyl)methyl]-5-methoxy-α-oxo- (CA INDEX NAME)

RN 260265-59-6 CAPLUS
CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-5-methoxy-α-οxo1-propyl- (CA INDEX NAME)

REFERENCE COUNT:

15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

CAPLUS COPYRIGHT 2008 ACS on STN ANSWER 4 OF 14

ACCESSION NUMBER:

1999:647583 CAPLUS

DOCUMENT NUMBER:

132:145941

TITLE:

Therapeutic potential of phosphodiesterase 4

inhibitors in allergic diseases

AUTHOR (S):

Crocker, I. Caroline; Townley, Robert G.

CORPORATE SOURCE:

Creighton University Allergic Disease Center, Omaha,

NE, USA

SOURCE:

Drugs of Today (1999), 35(7), 519-535 CODEN: MDACAP; ISSN: 0025-7656

PUBLISHER:

Prous Science

DOCUMENT TYPE:

Journal; General Review

LANGUAGE:

English

A review with 137 refs. CAMP is thought to be associated with inflammatory cell activity: high levels tend to decrease proliferation and cytokine secretion, whereas low concns. have the opposite effect (1). Since many phosphodiesterases (PDEs) degrade cAMP, inhibitors of this enzyme decrease inflammatory cell activity. Theophylline, which has nonselective PDE inhibitor activity in addition to its other mechanisms of action, has been used in the treatment of asthma for many years. Unfortunately, because of the important role of PDEs in the cell, nonspecific inhibition of these enzymes causes many undesirable side effects. The discovery of PDE isoenzyme families (PDE1-PDE10), their subtypes (HPDE4 and LPDE4) and their differential distribution among the cell types, as well as their specific functions in controlling cell processes, has led to the development of new, specific PDE4 inhibitors. This review details the rationale for the use of PDE4 inhibitors in the treatment of allergic disease. In addition, the effects of PDE4 inhibitors in vitro, in preclin. animal models and in the clinic are covered. Finally, up-to-date information on the most recently developed inhibitors, such as SB-207499, CDP-840, AWD-12-281 and D-4418, is provided.

IT 257892-33-4, AWD 12-281

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(therapeutic potential of phosphodiesterase 4 inhibitors in allergic diseases)

RN 257892-33-4 CAPLUS

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-CN fluorophenyl)methyl]-5-hydroxy-α-oxo- (CA INDEX NAME)

REFERENCE COUNT:

137 THERE ARE 137 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE REFORMAT

L8 ANSWER 5 OF 14 WPIDS COPYRIGHT 2008 THE THOMSON CORP ON STN DATA NOT AVAILABLE FOR THIS ACCESSION NUMBER

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L8 ANSWER 13 OF 14 WPIDS COPYRIGHT 2008 THE THOMSON CORP ON STN DATA NOT AVAILABLE FOR THIS ACCESSION NUMBER

L8 ANSWER 14 OF 14 USPATFULL on STN

ACCESSION NUMBER:

INVENTOR (S):

1999:170623 USPATFULL

TITLE:

N-substituted indole-3 glyoxylamides having

anti-asthmatic antiallergic and

immunosuppressant/immuno-modulating action

Lebaut, Guillaume, Saint Sebastien/Loire, France

Menciu, Cecilia, Nantes, France

Kutscher, Bernhard, Maintal, Germany, Federal Republic

of

Emig, Peter, Bruchkobel, Germany, Federal Republic of Szelenyi, Stefan, Schwaig, Germany, Federal Republic of Brune, Kay, Marloffstein/Rathsberg, Germany, Federal

Republic of

PATENT ASSIGNEE(S):

ASTA Medica Aktiengesellschgt, Germany, Federal

Republic of (non-U.S. corporation)

NUMBER KIND DATE PATENT INFORMATION: US 6008231 19991228 19970908 (8) APPLICATION INFO.: US 1997-925326 NUMBER PRIORITY INFORMATION: DE 1996-19636150 DOCUMENT TYPE: Utility FILE SEGMENT: Granted Richter, Johann PRIMARY EXAMINER: Oswecki, Jane C. ASSISTANT EXAMINER: LEGAL REPRESENTATIVE: Pillsbury Madison & Sutro NUMBER OF CLAIMS: 11 EXEMPLARY CLAIM: 1 942 LINE COUNT: CAS INDEXING IS AVAILABLE FOR THIS PATENT. The invention relates to novel N-substituted indole-3-glyoxylamides, to AΒ processes for their preparation and to their pharmaceutical use. The compounds have antiasthmatic, antiallergic and immunosuppressant/immunomodulating actions. CAS INDEXING IS AVAILABLE FOR THIS PATENT. IT 204205-78-7P 204205-79-8P 204205-80-1P 204205-81-2P 204205-82-3P 204205-85-6P 204205-86-7P 204205-87-8P 204205-90-3P 204205-91-4P 204205-92-5P 204205-93-6P 204205-95-8P 204205-96-9P 204205-97-0P 204205-98-1P 204206-01-9P 204206-02-0P 204206-03-1P (preparation of N-substituted indoleglyoxylamides as antiasthmatics, antiallergic agents and immunosuppressants/immunomodulators)

204205-78-7 USPATFULL

pyridinyl- (CA INDEX NAME)

RN

CN

1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]- α -oxo-N-4-

RN 204205-80-1 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]- α -oxo-N-3-pyridinyl- (CA INDEX NAME)

RN 204205-81-2 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-1-(phenylmethyl)-N-3-pyridinyl- (CA INDEX NAME)

RN 204205-85-6 USPATFULL CN 1H-Indole-3-acetamide, N-(2-chloro-3-pyridinyl)-1-[(4-fluorophenyl)methyl]- α -oxo- (CA INDEX NAME)

RN 204205-86-7 USPATFULL CN 1H-Indole-3-acetamide, α-oxo-1-(phenylmethyl)-N-4-pyridinyl- (CA INDEX NAME)

RN 204205-87-8 USPATFULL CN 1H-Indole-3-acetamide, α -oxo-N-4-pyridinyl-1-(3-pyridinylmethyl)- (CA INDEX NAME)

RN 204205-90-3 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 204205-91-4 USPATFULL CN 1H-Indole-3-acetamide, 1-[(2-chlorophenyl)methyl]- α -oxo-N-4-pyridinyl- (CA INDEX NAME)

RN 204205-92-5 USPATFULL CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]- α -oxo-N-2-pyridinyl- (CA INDEX NAME)

RN 204205-93-6 USPATFULL CN 1H-Indole-3-acetamide, α-oxo-N-4-pyridinyl-1-(2-pyridinylmethyl)-(CA INDEX NAME)

RN 204205-95-8 USPATFULL
CN 1H-Indole-3-acetamide, α-oxo-1-(phenylmethyl)-N-2-pyridinyl- (CF INDEX NAME)

RN 204205-96-9 USPATFULL CN Carbamic acid, [1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-6-yl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 204205-97-0 USPATFULL CN Carbamic acid, [1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-5-yl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 204205-98-1 USPATFULL

CN Carbamic acid, [1-[(4-fluorophenyl)methyl]-3-[oxo(4-pyridinylamino)acetyl]-1H-indol-6-yl]-, cyclopentyl ester (9CI) (CA INDEX NAME)

RN 204206-01-9 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-5-methoxy-α-οxο-N-4-pyridinyl- (CA INDEX NAME)

RN 204206-02-0 USPATFULL

CN 1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-5-hydroxy-α-οxο-N-4-pyridinyl- (CA INDEX NAME)

RN 204206-03-1 USPATFULL
CN Carbamic acid, [[1-[(4-fluorophenyl)methyl]-3-[oxo(4pyridinylamino)acetyl]-1H-indol-5-yl]methyl]-, ethyl ester (9CI) (CA
INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & &$$

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FILE CONTAINS CURRENT INFORMATION.
LAST RELOADED: Jan 11, 2008 (20080111/UP).

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(FILE 'HOME' ENTERED AT 15:23:09 ON 24 JAN 2008)

FILE 'REGISTRY' ENTERED AT 15:23:20 ON 24 JAN 2008

L1 STRUCTURE UPLOADED

L2 143 S L1 FULL

FILE 'MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 15:23:58 ON 24 JAN

| | 2008 | | | | | |
|----|------|-----|---|----|-----|---------------------------|
| L3 | | 185 | S | L2 | | |
| L4 | | 58 | S | L3 | AND | (CANCER? OR TUMOR?) |
| L5 | | | _ | | | ANGIOGENESIS |
| L6 | | 15 | S | L4 | AND | (MULTIDRUG OR MULTI-DRUG) |
| L7 | | 1 | S | L4 | NOT | PY>2000 |
| L8 | | 14 | S | L3 | NOT | PY>2000 |
| | | | | | | |

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